

10/009317

FILE 'REGISTRY' ENTERED AT 12:11:48 ON 27 OCT 2004  
L17 7 S GFCRCICTRGFCRCICTR | GVCRCLCRRGVCRCLCRR/SQSP

FILE 'CAPLUS' ENTERED AT 12:12:58 ON 27 OCT 2004  
L18 5 S L17

L18 ANSWER 1 OF 5 CAPLUS COPYRIGHT 2004 ACS on STN  
ED Entered STN: 26 Jan 2004  
ACCESSION NUMBER: 2004:60123 CAPLUS  
DOCUMENT NUMBER: 140:122752  
TITLE: Antimicrobial theta defensins, analogs thereof, and  
methods of use  
INVENTOR(S): Selsted, Michael E.; Tran, Dat Q.  
PATENT ASSIGNEE(S): The Regents of the University of California, A  
California Corporation, USA  
SOURCE: U.S. Pat. Appl. Publ., 46 pp.  
CODEN: USXXCO  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2004014669	A1	20040122	US 2003-427715	20030430
PRIORITY APPLN. INFO.:			US 2002-377071P	P 20020430
OTHER SOURCE(S):	MARPAT 140:122752			

AB The invention provides theta defensin analogs having antimicrobial activity. The invention also provides a method of reducing or inhibiting growth or survival of a microorganism in an environment capable of sustaining the growth or survival of the microorganism, comprising administering an effective amount of a theta defensin analog to the environment, thereby reducing or inhibiting the growth or survival of the microorganism. The structure and microbicidal activities and relationships of theta defensins and protegrin-1 were evaluated by comparing the microbicidal activities of 20 analogs against Escherichia coli, Candida albicans, and Cryptococcus neoformans and by determining the relative bactericidal activities in assays containing ionic and serum additives.

IT 306966-04-1P 374088-87-6P 648858-22-4P  
648858-23-5P 648858-24-6P  
RL: BSU (Biological study, unclassified); PAC (Pharmacological activity);  
PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL  
(Biological study); PREP (Preparation); USES (Uses)  
(antimicrobial theta defensins, analogs thereof, and uses)

L18 ANSWER 2 OF 5 CAPLUS COPYRIGHT 2004 ACS on STN  
ED Entered STN: 09 Aug 2002  
ACCESSION NUMBER: 2002:594692 CAPLUS  
DOCUMENT NUMBER: 137:153832  
TITLE: Novel antiviral activities of primate theta defensins  
and mammalian cathelicidins  
INVENTOR(S): Maury, Wendy; Stapleton, Jack; Stinski, Mark; Roller,  
Richard; McCray, Paul B.; Tack, Brian  
PATENT ASSIGNEE(S): University of Iowa Research Foundation, USA  
SOURCE: PCT Int. Appl., 65 pp.

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DOCUMENT TYPE: CODEN: PIXXD2  
 LANGUAGE: Patent  
 FAMILY ACC. NUM. COUNT: English  
 PATENT INFORMATION: 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002060468	A2	20020808	WO 2002-US2435	20020129
WO 2002060468	A3	20030123		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
US 2003022829	A1	20030130	US 2002-60102	20020129
US 2004086535	A1	20040506	US 2003-721839	20031125
PRIORITY APPLN. INFO.:			US 2001-265270P	P 20010130
			US 2001-309368P	P 20010801
			US 2002-60102	A3 20020129
AB The present invention relates to the use of anti-viral peptides in the inhibition and treatment of viral infections, in particular infections caused by enveloped viruses. These anti-viral peptides, some natural and others artificial, adopt either amphiphilic alpha-helical or a theta structure where the homodimeric or heterodimer peptides are joined by both cysteine bonds and circularization of the peptides. These agents may be used alone or in combination with more traditional anti-viral pharmaceuticals.				
IT 307334-75-4 307334-76-5				
RL: PRP (Properties)				
(unclaimed sequence; novel antiviral activities of primate theta defensins and mammalian cathelicidins)				
L18 ANSWER 3 OF 5 CAPLUS COPYRIGHT 2004 ACS on STN				
ED Entered STN: 07 Feb 2002				
ACCESSION NUMBER: 2002:102701 CAPLUS				
DOCUMENT NUMBER: 136:400525				
TITLE: Homodimeric $\theta$ -defensins from Rhesus macaque leukocytes. Isolation, synthesis, antimicrobial activities, and bacterial binding properties of the cyclic peptides				
AUTHOR(S): Tran, Dat; Tran, Patti A.; Tang, Yi-Quan; Yuan, Jun; Cole, Tim; Selsted, Michael E.				
CORPORATE SOURCE: Departments of Pathology and Microbiology & Molecular Genetics, University of California, Irvine, CA, 92697, USA				
SOURCE: Journal of Biological Chemistry (2002), 277(5), 3079-3084				
CODEN: JBCHA3; ISSN: 0021-9258				
PUBLISHER: American Society for Biochemistry and Molecular Biology				

Searcher : Shears 571-272-2528

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DOCUMENT TYPE: Journal  
LANGUAGE: English

AB Rhesus  $\theta$ -defensin 1 (RTD-1) is a unique tridisulfide, cyclic antimicrobial peptide formed by the ligation of two 9-residue sequences derived from heterodimeric splicing of similar 76-amino acid,  $\alpha$ -defensin-related precursors, termed RTD1a and RTD1b. The structures of RTD-2 and RTD-3 were predicted to exist if homodimeric splicing of the RTD1a and RTD1b occurs in vivo. Western blotting disclosed the presence of putative  $\theta$ -defensins, distinct from RTD-1, in leukocyte exts. Two new  $\theta$ -defensins, RTD-2 and RTD-3, were purified by reverse-phase high performance liquid chromatog. and characterized by amino acid anal., matrix-assisted laser desorption/ionization time-of-flight mass spectroscopy, and comparison to the synthetic stds. RTD-2 and RTD-3 are the predicted homodimeric splicing products of RTD1b and RTD1a, resp. The cellular abundance of RTD-1, -2, and -3 were 29:1:2, indicating that there is a preference for the heterodimeric ligation that generates RTD-1. RTD-1, -2, and -3 had similar antimicrobial activities against *Staphylococcus aureus*, *Candida albicans*, and *Cryptococcus neoformans*, whereas the activity of RTD-2 against *Escherichia coli* was 2-3-fold less than those of RTD-1 and RTD-3. Equal amts. of each  $\theta$ -defensin bound to *E. coli* cells, indicating that the differences in antibacterial activities are the result of post-binding processes.

IT 306966-04-1P 374088-87-6P

RL: BSU (Biological study, unclassified); PRP (Properties); PUR (Purification or recovery); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(isolation, synthesis, and antimicrobial activities of homodimeric  $\theta$ -defensins of Rhesus macaque)

REFERENCE COUNT: 45 THERE ARE 45 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L18 ANSWER 4 OF 5 CAPLUS COPYRIGHT 2004 ACS on STN

ED Entered STN: 20 Sep 2001

ACCESSION NUMBER: 2001:688259 CAPLUS

DOCUMENT NUMBER: 135:370575

TITLE: Circular minidefensins and posttranslational generation of molecular diversity

AUTHOR(S): Leonova, Larisa; Kokryakov, Vladimir N.; Aleshina, Galina; Hong, Teresa; Nguyen, Tung; Zhao, Chengquan; Waring, Alan J.; Lehrer, Robert I.

CORPORATE SOURCE: Department of Medicine, UCLA School of Medicine, Los Angeles, CA, USA

SOURCE: Journal of Leukocyte Biology (2001), 70(3), 461-464  
CODEN: JLBIE7; ISSN: 0741-5400

PUBLISHER: Federation of American Societies for Experimental Biology

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The authors purified two new minidefensins (RTD-2 and RTD-3) from the bone marrow of rhesus monkeys. Both were circular octadecapeptides that contained three intramol. disulfide bonds and were homologous to RTD-1, a circular ( $\theta$ ) defensin described previously. However, whereas the 18 residues of RTD-1 represent spliced nonapeptide fragments derived from two different demidefensin precursors, RTD-2 and -3 comprise tandem nonapeptide repeats derived from only one of the RTD-1 precursors. Thus,

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circular minidefensins are products of a novel post-translational system that generates effector mol. diversity without commensurate genome expansion. A system wherein two demidefensin genes can produce three circular minidefensins might allow n such genes to produce (n/2)(n+1) peptides.

IT 306966-04-1,  $\theta$ -Defensin RTD 3 374088-87-6,  
 $\theta$ -Defensin RTD 2

RL: BAC (Biological activity or effector, except adverse); BOC (Biological occurrence); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study); OCCU (Occurrence)

(cloning and characterization of circular defensins of rhesus monkey)

REFERENCE COUNT: 10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L18 ANSWER 5 OF 5 CAPLUS COPYRIGHT 2004 ACS on STN

ED Entered STN: 21 Nov 2000

ACCESSION NUMBER: 2000:814517 CAPLUS

DOCUMENT NUMBER: 133:366399

TITLE: Antimicrobial theta-defensins and methods of using same

INVENTOR(S): Selsted, Michael E.; Tang, Yi-quan; Yuan, Jun; Ouellette, Andre J.

PATENT ASSIGNEE(S): The Regents of the University of California, USA

SOURCE: PCT Int. Appl., 110 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000068265	A1	20001116	WO 2000-US12842	20000510
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
US 6335318	B1	20020101	US 1999-309487	19990510
EP 1187850	A1	20020320	EP 2000-930577	20000510
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO			
US 6514727	B1	20030204	US 2001-967808	20010926
US 2003162718	A1	20030828	US 2002-313994	20021205
PRIORITY APPLN. INFO.:			US 1999-309487	A2 19990510
			WO 2000-US12842	W 20000510
			US 2001-967808	A1 20010926

OTHER SOURCE(S): MARPAT 133:366399

AB The present invention relates to an isolated cyclic peptide,  $\theta$ -defensin, having antimicrobial activity, and to  $\theta$ -defensin analogs. A  $\theta$ -defensin can have the amino acid sequence

Searcher : Shears 571-272-2528

Xaa1-Xaa2-Xaa3-Xaa4-Xaa5-Xaa1-Xaa6-Xaa4-Xaa4-Xaa1-Xaa1-Xaa6-Xaa4-Xaa5-Xaa1-Xaa3-aa7-Xaa8, wherein Xaa1 to Xaa8 are defined; wherein Xaa1 can be linked through a peptide bond to Xaa8; and wherein crosslinks can be formed between Xaa3 and Xaa3, between Xaa5 and Xaa5, and between Xaa7 and Xaa7. For example, the invention provides a  $\theta$ -defensin having the amino acid sequence Gly-Phe-Cys-Arg-Cys-Leu-Cys-Arg-Arg-Gly-Val-Cys-Arg-Cys-Ile-Cys-Thr-Arg (SEQ ID NO:1), wherein the Gly at position 1 (Gly-1) is linked through a peptide bond to Arg-18, and wherein disulfide bonds are present between Cys-3 and Cys-16, between Cys-5 and Cys-14, and between Cys-7 and Cys-12. The invention also provides nucleic acids encoding  $\theta$ -defensins and antibodies that specifically bind a  $\theta$ -defensin. In addition, the invention relates to methods of using  $\theta$ -defensin to reduce or inhibit microbial growth or survival.

IT **306966-04-1P**

RL: BAC (Biological activity or effector, except adverse); BPN (Biosynthetic preparation); BSU (Biological study, unclassified); PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
(amino acid sequence; antimicrobial theta-defensins and methods of using same)

IT **307334-75-4 307334-76-5**

RL: PRP (Properties)  
(unclaimed sequence; antimicrobial theta-defensins and methods of using same)

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

E20 THROUGH E26 ASSIGNED

FILE 'REGISTRY' ENTERED AT 12:13:42 ON 27 OCT 2004

L19 7 SEA FILE=REGISTRY ABB=ON PLU=ON (306966-04-1/BI OR 374088-87-6/BI OR 307334-75-4/BI OR 307334-76-5/BI OR 648858-22-4/BI OR 648858-23-5/BI OR 648858-24-6/BI)

L20 7 L17 AND L19

L20 ANSWER 1 OF 7 REGISTRY COPYRIGHT 2004 ACS on STN

RN **648858-24-6** REGISTRY

CN L-Argininamide, glycyl-L-valyl-L-cysteinyl-L-arginyl-L-cysteinyl-L-leucyl-L-cysteinyl-L-arginyl-L-arginylglycyl-L-valyl-L-cysteinyl-L-arginyl-L-cysteinyl-L-leucyl-L-cysteinyl-L-arginyl-, cyclic (3→16), (5→14), (7→12)-tris(disulfide) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 7: PN: US20040014669 TABLE: 1 claimed protein

SQL 18

SEQ 1 GVCRCICRRG VCRCLCRR

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HITS AT: 1-18

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 140:122752

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L20 ANSWER 2 OF 7 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 648858-23-5 REGISTRY  
CN L-Arginine, glycyl-L-valyl-L-cysteinyl-L-arginyl-L-cysteinyl-L-leucyl-L-cysteinyl-L-arginyl-L-arginylglycyl-L-valyl-L-cysteinyl-L-arginyl-L-cysteinyl-L-leucyl-L-cysteinyl-L-arginyl-, cyclic (3→16), (5→14), (7→12)-tris(disulfide) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 6: PN: US20040014669 TABLE: 1 claimed protein  
SQL 18

SEQ 1 GVCRLCRRG VCRCLCRR  
=====

HITS AT: 1-18

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 140:122752

L20 ANSWER 3 OF 7 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 648858-22-4 REGISTRY  
CN L-Arginine, glycyl-L-phenylalanyl-L-cysteinyl-L-arginyl-L-cysteinyl-L-isoleucyl-L-cysteinyl-L-threonyl-L-arginylglycyl-L-phenylalanyl-L-cysteinyl-L-arginyl-L-cysteinyl-L-isoleucyl-L-cysteinyl-L-threonyl-, cyclic (3→16), (5→14), (7→12)-tris(disulfide) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 5: PN: US20040014669 TABLE: 1 claimed protein  
SQL 18

SEQ 1 GFCRCICTRG FCRCICTR  
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HITS AT: 1-18

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 140:122752

L20 ANSWER 4 OF 7 REGISTRY COPYRIGHT 2004 ACS on STN  
RN 374088-87-6 REGISTRY  
CN Cyclo(L-arginyl-L-arginylglycyl-L-valyl-L-cysteinyl-L-arginyl-L-cysteinyl-L-leucyl-L-cysteinyl-L-arginyl-L-arginylglycyl-L-valyl-L-cysteinyl-L-arginyl-L-cysteinyl-L-leucyl-L-cysteinyl), cyclic (5→18), (7→16), (9→14)-tris(disulfide) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 0-Defensin RTD 2  
CN 3: PN: US20040014669 TABLE: 1 claimed protein  
SQL 18

SEQ 1 RRGVCRCLCR RGVCRCLC  
=====

HITS AT: 1-11, 3-18

REFERENCE 1: 140:122752

10/009317

REFERENCE 2: 136:400525

REFERENCE 3: 135:370575

L20 ANSWER 5 OF 7 REGISTRY COPYRIGHT 2004 ACS on STN

RN 307334-76-5 REGISTRY

CN L-Arginine, glycyl-L-valyl-L-cysteinyl-L-arginyl-L-cysteinyl-L-leucyl-L-cysteinyl-L-arginyl-L-arginylglycyl-L-valyl-L-cysteinyl-L-arginyl-L-cysteinyl-L-leucyl-L-cysteinyl-L-arginyl- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 12: PN: WO0068265 FIGURE: 16 unclaimed sequence

CN 29: PN: WO02060468 SEQID: 29 unclaimed sequence

SQL 18

SEQ 1 GVCRCLCRRG VCRCLCRR.

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HITS AT: 1-18

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 137:153832

REFERENCE 2: 133:366399

L20 ANSWER 6 OF 7 REGISTRY COPYRIGHT 2004 ACS on STN

RN 307334-75-4 REGISTRY

CN L-Arginine, glycyl-L-phenylalanyl-L-cysteinyl-L-arginyl-L-cysteinyl-L-isoleucyl-L-cysteinyl-L-threonyl-L-arginylglycyl-L-phenylalanyl-L-cysteinyl-L-arginyl-L-cysteinyl-L-isoleucyl-L-cysteinyl-L-threonyl- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 11: PN: WO0068265 FIGURE: 16 unclaimed sequence

CN 28: PN: WO02060468 SEQID: 28 unclaimed sequence

SQL 18

SEQ 1 GFCRCICTRG FCRCICTR

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HITS AT: 1-18

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

REFERENCE 1: 137:153832

REFERENCE 2: 133:366399

L20 ANSWER 7 OF 7 REGISTRY COPYRIGHT 2004 ACS on STN

RN 306966-04-1 REGISTRY

CN Cyclo(L-arginyl-L-cysteinyl-L-isoleucyl-L-cysteinyl-L-threonyl-L-arginylglycyl-L-phenylalanyl-L-cysteinyl-L-arginyl-L-cysteinyl-L-isoleucyl-L-cysteinyl-L-threonyl-L-arginylglycyl-L-phenylalanyl-L-cysteinyl), cyclic (2→11), (4→9), (13→18)-tris(disulfide) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 0-Defensin RTD 3

CN 2: PN: US20040014669 TABLE: 1 claimed protein

SQL 18

10/009317

SEQ 1 RCICTRGFCR CICTRGFC

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HITS AT: 1-15, 7-18

REFERENCE 1: 140:122752

REFERENCE 2: 136:400525

REFERENCE 3: 135:370575

REFERENCE 4: 133:366399

(FILE 'MEDLINE, BIOSIS, EMBASE' ENTERED AT 12:15:10 ON 27 OCT 2004)  
L22 0 S L17

FILE 'HOME' ENTERED AT 12:15:19 ON 27 OCT 2004